

<b>TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US) CONCERNING A SUBMISSION UNDER 35 U.S.C. 371</b>		ATTORNEY'S DOCKET NUMBER 19672-004US1 RET/PCG-9002US U.S. APPLICATION NO. (if known, see 37 C.F.R. 1.5) <div style="font-size: 2em; font-weight: bold; text-align: center;">10/570647</div>
INTERNATIONAL APPLICATION NO. PCT/JP2004/013183	INTERNATIONAL FILING DATE 3 September 2004	PRIORITY DATE CLAIMED 4 September 2003
TITLE OF INVENTION REMEDY AND DETECTION DRUG FOR BILE DUCT CANCER		
APPLICANT(S) FOR DO/EO/US Hiroyuki Aburatani, Masahisa Fukayama, Naoko Yamauchi, Masami Suzuki, Atsuhiko Kato and Masayuki Tsuchiya		

Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:

1. ☒ This is a **FIRST** submission of items concerning a submission under 35 U.S.C. 371.
2. ☐ This is a **SECOND** or **SUBSEQUENT** submission of items concerning a submission under 35 U.S.C. 371.
3. ☐ This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below.
4. ☒ The US has been elected (Article 31).
5. ☒ A copy of the International Application as filed (35 U.S.C. 371(c)(2))
  - a. ☐ is attached hereto (required only if not communicated by the International Bureau).
  - b. ☒ has been communicated by the International Bureau.
  - c. ☐ is not required, as the application was filed in the United States Receiving Office (RO/US).
6. ☒ An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)).
  - a. ☒ is attached hereto.
  - b. ☐ has been previously submitted under 35 U.S.C. 154(d)(4).
7. ☒ Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))
  - a. ☐ are attached hereto (required only if not communicated by the International Bureau).
  - b. ☐ have been communicated by the International Bureau.
  - c. ☐ have not been made; however, the time limit for making such amendments has NOT expired.
  - d. ☒ have not been made and will not be made.
8. ☐ An English language translation of amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)).
9. ☐ An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).
10. ☐ An English language translation of the annexes to the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).

**Items 11 to 20 below concern document(s) or information included:**

11. ☐ An Information Disclosure Statement under 37 C.F.R. 1.97 and 1.98.
12. ☐ An assignment document for recording. A separate cover sheet in compliance with 37 C.F.R. 3.28 and 3.31 is included.
13. ☐ A preliminary amendment.
14. ☐ An Application Data Sheet under 37 C.F.R. 1.76.
15. ☐ A substitute specification.
16. ☐ A power of attorney and/or change of address letter.
17. ☐ A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 37 C.F.R. 1.821 - 1.825.
18. ☐ A second copy of the published international application under 35 U.S.C. 154(d)(4).
19. ☐ A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).

**CERTIFICATE OF MAILING BY EXPRESS MAIL**  
 Express Mail Label No. EV 664070748 US  
 Date of Deposit: March 3, 2006

U.S. APPLICATION NO. (if known, see 37 C.F.R. 1.5) <b>10/570647</b>		INTERNATIONAL APPLICATION NO. PCT/JP2004/013183		ATTORNEY'S DOCKET NUMBER 19672-004US1	
20. Other items or information: International Search Report Submission of Substitute Specification Substitute specification (version with markings to show changes made) Substitute specification (clean copy)					
The following fees have been submitted				<b>CALCULATIONS PTO USE ONLY</b>	
21. <input checked="" type="checkbox"/> Basic national fee (37 CFR 1.192(a)).....\$300				\$300.00	
22. <input checked="" type="checkbox"/> Examination fee (37 CFR 1.492(c)) If the written opinion prepared by ISA/US or the international preliminary examination report prepared by the IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4) .....\$0 All other situations .....\$200				\$200.00	
23. <input checked="" type="checkbox"/> Search fee (37 CFR 1.92(b)) If the written opinion of the ISA/US or the international preliminary examination report prepared by IPEA/US indicates all claims satisfy provisions of PCT Article 33(1)-(4) .....\$0 Search fee (37 CFR 1.445(a)(2)) has been paid on the international application to the USPTO as an International Searching Authority.....\$100 International Search Report prepared by an ISA other than the US and provided to the Office or previously communicated to the US by the IB .....\$400 All other situations .....\$500				\$400.00	
<b>Total of 21, 22 and 23 =</b>				\$900.00	
<input type="checkbox"/> Additional fee for specification and drawings filed in paper over 100 sheets (excluding sequence listing in compliance with 37 CFR 1.821(c) or (e) or computer program listing filed in an electronic medium) (37 CFR 1.492(j)). The fee is \$250 for each additional 50 sheets of paper or fraction thereof.					
Total Sheets	Extra Sheets	Number of each additional 50 or fraction thereof (round up to a whole number)	RATE		
-100 =	/50=		X \$250	\$0.00	
Surcharge of \$130 for furnishing any of the search fee, examination fee, or the oath or declaration after the date of commencement of the national stage (37 C.F.R. 1.492(h)).				\$0.00	
CLAIMS	NUMBER FILED	NUMBER EXTRA	RATE		
Total Claims	20 - 20 =	0	x \$50	\$0.00	
Independent Claims	5 - 3 =	2	x \$200	\$400.00	
MULTIPLE DEPENDENT CLAIM(S) (if applicable)			+ \$360	\$360.00	
<b>TOTAL OF ABOVE CALCULATIONS =</b>				\$1,660.00	
<input type="checkbox"/> Applicant claims small entity status. See 37 C.F.R. 1.27. Fees above are reduced by 1/2.				\$0.00	
<b>SUBTOTAL =</b>				\$1,660.00	
Processing fee of \$130 for furnishing the English translation later than 30 months from the earliest claimed priority date (37 C.F.R. 1.492(i))				\$0.00	
<b>TOTAL NATIONAL FEE =</b>				\$1,660.00	
Fee for recording the enclosed assignment (37 C.F.R. 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 C.F.R. 3.28, 3.31). \$40.00 per property				\$0.00	
<b>TOTAL FEES ENCLOSED =</b>				\$1,660.00	
				<b>Amount to be refunded:</b>	\$0.00
				<b>Amount to be charged:</b>	\$0.00

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REGISTERED 03 MAR 2006

SUBSTITUTE PTO-1390

- a. ☒ A check in the amount of \$1,660.00 to cover the above fees is enclosed.
- b. ☐ Please charge my Deposit Account No. 06-1050 in the amount of \$0.00 to cover the above fees.  
A duplicate copy of this sheet is enclosed.
- c. ☒ The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 06-1050. A duplicate copy of this sheet is enclosed.
- d. ☐ Fees are to be charged to a credit card. **WARNING:** Information on this form may become public. **Credit card information should not be included on this form.** Provide credit card information and authorization on PTO-2038.

**NOTE:** Where an appropriate time limit under 37 C.F.R. 1.495 has not been met, a petition to revive (37 C.F.R. 1.137(a) or (b) must be filed and granted to restore the International Application to pending status.

SEND ALL CORRESPONDENCE TO:

PTO Customer No: 26161

SIGNATURE

Janis K. Fraser, Ph.D., J.D.

NAME

34,819

REGISTRATION NUMBER

21279172.doc

10/573647

Attorney's Docket No.: 19672-004US1 / RET/PCG-9002US

1453 Rec'd PGT/770 03 MAR 2006

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant : Hiroyuki Aburatani et al.      Art Unit : Unknown  
Serial No. : To be Assigned      Examiner : Unknown  
Filed : Herewith  
Title : REMEDY AND DETECTION DRUG FOR BILE DUCT CANCER

MAIL STOP PCT  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

STATEMENT OF SUBSTITUTE SPECIFICATION UNDER 37 CFR § 1.125

Pursuant to 37 CFR § 1.125, Applicants submit a substitute specification encompassing changes being made to the specification filed herewith. No new matter has been introduced by this substitution to the specification filed. Applicants request entry of the substitute specification.

Also attached is a marked-up version showing the changes that have been made.

Please apply any charges or credits to Deposit Account No. 06-1050.

Respectfully submitted,

Date: March 3, 2006

John W. Freeman Reg No. 29,066  
Janis K. Fraser, Ph.D., J.D.  
Reg. No. 34,819

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21279771.doc

CERTIFICATE OF MAILING BY EXPRESS MAIL

Express Mail Label No. EV 664070748 US

March 3, 2006  
Date of Deposit